Substitute for PTO-1390 (Rev. 07-2005) "Transmittal Letter to the United States Approved for use through 3/31/2007. OMB 0651-0021 ATTORNEY'S DOCKET NUMBER U. S. PATENT AND TRADEMARK OFFICE U.S. DEPARTMENT OF COMMERCE ITR0041P TRANSMITTAL LETTER TO THE UNITED STATES U.S. APPLICATION NO. (If known, see 37 CFR 1.5) DESIGNATED/ELECTED OFFICE (DO/EO/US) **CONCERNING A FILING UNDER 35 U.S.C. 371** INTERNATIONAL APPLICATION NO. INTERNATIONAL FILING DATE PRIORITY DATE CLAIMED PCT/EP2004/011683 15 October 2004 24 October 2003 TITLE OF INVENTION ORTHOGONAL GENE SWITCHES APPLICANT(S) FOR DO/EO/US Gennaro Ciliberto, Raffaele De Francesco, Daniela Fattori, Paola Gallinari, Olaf D. Kinzel, Uwe Koch, Ester Muraglia, Carlo Toniatti, Riccardo Cortese and Armin Lahm Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information: 1. This is a **FIRST** submission of items concerning a filing under 35 U.S.C. 371. This is a SECOND or SUBSEQUENT submission of items concerning a filing under 35 U.S.C. 371. This is an express request to begin national examination procedures [35 U.S.C. 371(f)]. The submission must include items (5), (6), (9) and (21) indicated below. The US has been elected (PCT Article 31). A copy of the International Application as filed [35 U.S.C. 371(c)(2)] a. is attached hereto (required only if not communicated by the International Bureau). b. X has been communicated by the International Bureau. c. is not required, as the application was filed in the United States Receiving Office (RO/US). An English language translation of the International Application as filed [35 U.S.C. 371(c)(2)]. a. is attached hereto b. has been previously submitted under 35 U.S.C. 154(d)(4). 7. Amendments to the claims of the International Application under PCT Article 19 [35 U.S.C. 371(c)(3)]. are attached hereto (required only if not communicated by the International Bureau). have been communicated by the International Bureau. c. have not been made; however, the time limit for making such amendments has NOT expired. d. have not been made and will not be made. An English language translation of the amendments to the claims under PCT Article 19 [35 U.S.C. 371(c)(3)]. An oath or declaration of the inventor(s) [35 U.S.C. 371(c)(4)]. ☐ An English language translation of the annexes of the International Preliminary Examination Report under PCT Article 36 [35 U.S.C. 371(c)(5)].

Items 11 to 20 below concern other document(s) or information included: An Information Disclosure Statement under 37 CFR 1.97 and 1.98. An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included. 13. A preliminary amendment. An Application Data Sheet under 37 CFR 1.76. 15. A substitute specification. 16. A power of attorney and/or change of address letter. 17. A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 CFR 1.821 - 1.825. 18. A second copy of the published International Application under 35 U.S.C.154(d)(4).

☐ A second copy of the English language translation of the international application under 35 U.S.C.154(d)(4).

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20. Other items or information:

IAP20 RCG'CLUTTO 24 APR 2006

Substitute for PTO-1390 (Rev. 07-2005) "Transmittal Letter to the United States Designated/Elected Office(DO/E sion Under 35 U.S.C. 371" Approved for use through 3/31/2007. OMB 0651-0021 INTERNATIONAL APPLICATION NO. U.S. APPLICATION NO. (If known see 37 CFR 15) ATTORNEY'S DOCKET NUMBER ITR0041P PCT/EP2004/011683 The following fees are submitted: CALCULATIONS PTO USE ONLY Basic national fee..... \$300.00 Examination fee (37 CFR 1.492(c)) If the written opinion prepared by ISA/US or the international preliminary examination report prepared by IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4)......\$0 \$200.00 Search fee (37 CFR 1.492(b)) If the written opinion of the ISA/US or the international preliminary examination report prepared by Search fee [37 CFR 1.445(a)(2)] has been paid on the international application to the USPTO as an International Searching Authority..... \$400.00 International Search Report prepared by an ISA other than the US and provided to the Office or previously communicated to the US by the IB TOTAL OF 21, 22 and 23= \$900.00 Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing or computer program listing filed in an electronic medium). The fee is \$250 for each additional 50 sheets of paper or fraction thereof. Number of each additional 50 or fraction **Total Sheets** Extra Sheets RATE thereof (round up to a whole number) 108 - 100 = /50 0.16 1 Х \$250.00 \$250.00 Surcharge of \$130.00 for furnishing the oath or declaration later than 30 months from the earliest claimed priority date [37 CFR 1.492(e)] CLAIMS NUMBER FILED NUMBER EXTRA RATE Total Claims - 20 = \$50.00 \$950.00 Х Independent Claims X - 3 = \$200.00 \$0.00 3 0 MULTIPLE DEPENDENT CLAIM(S) (if applicable) \$360.00 TOTAL OF ABOVE CALCULATIONS (SUBTOTAL) = \$2,100.00 Processing fee of \$130.00 for furnishing the English translation later than 30 months from the earliest claimed priority date (37 CFR 1.492(f)]. = TOTAL NATIONAL FEE \$2,100.00 Fee for recording the enclosed assignment [37 CFR 1.21(h)]. The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31).\$40.00 per property. TOTAL FEES ENCLOSED = \$2,100.00 Amount to be refunded: Amount to be charged: A check in the amount of \$ _ _ to cover the above fees is enclosed. Please charge my Deposit Account No. 13-2755 in the amount of \$2,100.00 A duplicate copy of this sheet is enclosed. The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to the Deposit Account No. 13-2755. A duplicate copy of this sheet is enclosed. NOTE: Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive [37 CFR 1.137(a) or (b)] must be filed and granted to restore the International Application to pending status. SEND ALL CORRESPONDENCE TO: Customer No. 000210 MERCK & CO., INC. Patent Department, RY60-30 P.O. Box 2000 126 East Lincoln Avenue Rahway, New Jersey 07065-0970 Heidi M. Struse **NAME** DATE: April 24, 2006 50,288

PHONE #: (732) 594-0238

REGISTRATION NUMBER

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Attorney Docket No.:	ITR0041P
International Appln No.:	PCT/EP2004/011683
International Filing Date:	15 October 2004
Priority Date:	24 October 2003
Title:	ORTHOGONAL GENE SWITCHES

Inventors:

Gennaro Ciliberto, Raffaele De Francesco, Daniela Fattori, Paola Gallinari, Olaf D. Kinzel, Uwe Koch, Ester Muraglia, Carlo Toniatti, Riccardo Cortese and Armin Lahm

EXPRESS MAIL CERTIFICATE

DATE OF DEPOSIT

April 24, 2006

EXPRESS MAIL NO. EVGLOSOSZO8345

I HEREBY CERTIFY THAT THIS CORRESPONDENCE IS BEING DEPOSITED WITH THE UNITED STATES POSTAL SERVICE AS EXPRESS MAIL "POST OFFICE TO ADDRESSEE" ON THE ABOVE DATE IN AN ENVELOPE ADDRESSED TO COMMISSIONER FOR PATENTS P.O. BOX 1450, ALEXANDRIA, VIRGINIA 22313-1450.

MAILED BY Loui Schejisi DATE 4-24-06